## **WEST Search History**

DATE: Thursday, May 22, 2003

	Hit Count	
		result set
PLUR=YES; OP=ADJ	r	
3667-\$.did.	2	L5
	0	L4
S; OP=ADJ		
ve t cell	0	L3
; OP=ADJ		
ve t cell	1	L2
ve t cells	1	L1
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END OF SEARCH HISTORY

FILE 'HOME' ENTERED AT 16:32:00 ON 22 MAY 2003

=> index bioscience

FILE 'DRUGMONOG' ACCESS NOT AUTHORIZED

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

INDEX 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOSIS, BIOTECHABS, BIOTECHDS, BIOTECHNO, CABA, CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN, CONFSCI, CROPB, CROPU, DDFB, DDFU, DGENE, DRUGB, DRUGLAUNCH, DRUGMONOG2, ...' ENTERED AT 16:32:07 ON 22 MAY 2003

67 FILES IN THE FILE LIST IN STNINDEX

Enter SET DETAIL ON to see search term postings or to view search error messages that display as 0\* with SET DETAIL OFF.

- => s (hyperactive t-cell)
  - 1 FILE BIOSIS
  - 1 FILE BIOTECHABS
  - 1 FILE BIOTECHDS
  - 1 FILE BIOTECHNO
  - 12 FILES SEARCHED...
    - 2 FILE CANCERLIT
    - 3 FILE CAPLUS
  - 23 FILES SEARCHED...
    - 3 FILE EMBASE
  - 32 FILES SEARCHED...
  - 44 FILES SEARCHED...
    - 3 FILE MEDLINE
    - 1 FILE PHARMAML
    - 1 FILE PHIN
    - 1 FILE PROMT
    - 2 FILE SCISEARCH
  - 60 FILES SEARCHED...
    - 2 FILE TOXCENTER
    - 2 FILE WPIDS
  - 66 FILES SEARCHED...
    - 2 FILE WPINDEX
  - 15 FILES HAVE ONE OR MORE ANSWERS, 67 FILES SEARCHED IN STNINDEX
- L1 QUE (HYPERACTIVE T-CELL)

=> d rank		
F1	3	CAPLUS
F2	3	<b>EMBASE</b>
F3	3	MEDLINE
F4	2	CANCERLIT
F5	2	SCISEARCH
F6	2	TOXCENTER
F7	2	WPIDS
F8	2	WPINDEX
F9	1	BIOSIS
F10	1	BIOTECHABS
F11	1	BIOTECHDS
F12	1	BIOTECHNO
F13	1	PHARMAML
F14	1	PHIN
F15	1	PROMT

=> file f1-15
COST IN U.S. DOLLARS

FULL ESTIMATED COST

SINCE FILE TOTAL ENTRY SESSION 2.20 2.41

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=> s l1

4 FILES SEARCHED...

8 FILES SEARCHED...

L2 23 L1

=> dup rem 12 DUPLICATE IS NOT AVAILABLE IN 'PHARMAML'. ANSWERS FROM THESE FILES WILL BE CONSIDERED UNIQUE PROCESSING COMPLETED FOR L2

11 DUP REM L2 (12 DUPLICATES REMOVED)
ANSWERS '1-3' FROM FILE CAPLUS
ANSWERS '4-5' FROM FILE EMBASE
ANSWER '6' FROM FILE TOXCENTER
ANSWERS '7-8' FROM FILE WPIDS
ANSWER '9' FROM FILE PHARMAML
ANSWER '10' FROM FILE PHIN
ANSWER '11' FROM FILE PROMT

```
ANSWER 1 OF 11 CAPLUS COPYRIGHT 2003 ACS
                                                     DUPLICATE 2
L3
     1999:27731 CAPLUS
ΔN
     130:76171
DN
    Use of proteolytic enzymes for treating glomerulonephritis
TI
IN
     Stauder, Gerhard; Ransberger, Karl
    Mucos Pharma G.m.b.H. & Co., Germany
PΑ
    PCT Int. Appl., 28 pp.
SO
    CODEN: PIXXD2
DT
    Patent
LA
    German
FAN.CNT 1
                                          APPLICATION NO. DATE
    PATENT NO.
                     KIND DATE
                                          -----
     ______
                           ------
                                          WO 1998-EP3769
PΙ
    WO 9858667
                     A1
                           19981230
                                                           19980619
        W: US
         RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
                                          DE 1997-19726253 19970620
    DE 19726253
                      A1
                           19981224
     DE 19726253
                      C2
                           20000316
     EP 920332
                      A1
                           19990609
                                          EP 1998-934986
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
PRAI DE 1997-19726253
                           19970620
     WO 1998-EP3769
                           19980619
     Proteinases, particularly trypsin, bromelain, and papain, are administered
AB
     for enzymic dissoln. of immune complexes deposited in the glomeruli in
     treatment of glomerulonephritis. The therapeutic effect of proteinases
     may also involve actions on cytokines, leukocyte cytokine receptors such
     as CD2, CD4, CD11b, and CD25, endogenous metalloproteinases, cell adhesion
     mols., and hyperactive T-cells.
             THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
    ANSWER 2 OF 11 CAPLUS COPYRIGHT 2003 ACS
                                                      DUPLICATE 4
L3
AN
     1987:494856 CAPLUS
DM
     107 - 94856
TI
     In vitro studies on leukemic cells and T lymphocytes in hairy cell
     leukemia
AU
     Ford, Richard J.; Mehta, Shashi; Sharma, Surendra
    M. D. Anderson Hosp. Tumor Inst., Univ. Texas, Houston, TX, 77030, USA
CS
    Leukemia (1987), 1(4), 386-9
SO
    CODEN: LEUKED; ISSN: 0887-6924
DT
     Journal
    English
LA
AB
    Hairy cell leukemia cell lines were established from 8 untreated patients
     using purified B cell growth factor (BCGF) in vitro. These cell lines
     maintained their original cell surface immunophenotype for about 1 mo,
     after which they began to lose 1 or more of their characteristic surface
               The cell lines also maintained typical hairy cell leukemia
     morphol. for 2-3 mo in vitro but later showed an increasing no. of
     multinucleate giant cells that maintained a B cell surface phenotype. The
     cell lines became independent of exogenously provided BCGF after at least
     1 mo in vitro and secreted BCGF activity into culture supernatants in most
     cases. Some cell lines also acquired Epstein-Barr virus nuclear antigen
    positivity after variable period. Two hairy cell leukemia patients also
     showed hyperactive T cell responses in vitro
     and exhibited spontaneous T cell proliferation in culture without
     exogenously supplied interleukin-2. These T cell lines had the T helper
```

phenotype and secreted significant amts. of T cell-assocd. lymphokines

with BCGF and interleukin-2 activity into culture supernatants.

L3

```
AN
     2000:260036 CAPLUS
DN
     132:274331
     Use of proteolytic enzymes to influence hyperactive T-
ΤI
IN
     Ransberger, Karl; Stauder, Gerhard
PA
     Mucos Pharma G.m.b.H. & Co., Germany
SO
     PCT Int. Appl., 27 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     German
FAN.CNT 1
     PATENT NO.
                      KIND DATE
                                          APPLICATION NO. DATE
                     ----
                                           -----
     WO 2000021547
                       A2
                            20000420
                                           WO 1999-EP7634
PΙ
                                                            19991012
                            20000908
     WO 2000021547
                      A3
         W: CA, US
         RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC,
             PT, SE
                            20000420
                                           DE 1998-19847114 19981013
     DE 19847114
                       A1
     EP 1121146
                                           EP 1999-953778 19991012
                       A2
                            20010808
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
PRAI DE 1998-19847114 A
                            19981013
     WO 1999-EP7634
                      W
                            19991012
AB
     The use of at least one proteolytic enzyme to influence
     hyperactive T cells is disclosed. Preferred
     proteolytic enzymes are trypsin, bromelain and papain. Rutin can addnl.
     be used.
     ANSWER 4 OF 11 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V.DUPLICATE 3
L3
     1998361153 EMBASE
AN
ΤI
     Immunopathogenesis of SLE.
     Mason L.J.; Isenberg D.A.
L.J. Mason, Bloomsbury Rheumatology Unit, Department of Medicine,
ΑU
CS
     University College London, 40-50 Tottenham Street, London W1P 9PG, United
     Kingdom
SO
     Bailliere's Clinical Rheumatology, (1998) 12/3 (385-403).
     Refs: 51
     ISSN: 0950-3579 CODEN: BCRHEZ
     United Kingdom
CY
DT
     Journal; Article
FS
     026
             Immunology, Serology and Transplantation
     031
             Arthritis and Rheumatism
     037
             Drug Literature Index
LA
     English
SL
     English
     Systemic lupus erythematosus (SLE) is an autoimmune rheumatic disease
     characterized by the deposition of autoantibodies and immune complexes,
     leading to tissue damage. The immunopathogenesis of SLE is like a jigsaw
     puzzle, some pieces of which are missing or have not fallen into place. In
    predisposed individuals, the initial stimulus is likely to be one or more
     of the environmental agents interacting with susceptibility genes. Once
     the critical threshold is breached there is a failure of the immune system
     to downregulate the ensuing abnormal immune response, involving polyclonal
    B cell activation and hyperactive T cell
    help. Key questions include, what are the processes behind the
     availability of autoantigens and the breakdown of tolerance that give rise
     to the pathogenic autoantibodies? Current areas of research also involve
     the roles played by cytokines, adhesion molecules, co-stimulatory
```

- L3 ANSWER 5 OF 11 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V.DUPLICATE 6
- AN 77039476 EMBASE
- DN 1977039476
- TI Hyperactive T cell function in young NZB

molecules and apoptosis.

mice; Increased proliferative responses to allogenic cells. ΑU Palmer D.W.; Dauphinee M.J.; Murphy E.; Talal N. CS Dept. Med., Univ. California, San Francisco, Calif., United States SO Clinical and Experimental Immunology, (1976) 23/3 (578-581). CODEN: CEXIAL Journal рΤ FS 026 Immunology, Serology and Transplantation 005 General Pathology and Pathological Anatomy LA English ÀΒ The one way mixed lymphocyte reaction was employed to study proliferative responses to antigens by mature, immunocompetent T cells from NZB mice 3 wk to 4 mth old. Compared to cells from control mice of the same H-2 type, thymus, spleen and lymph node cells from NZB mice were hyperactive in this response. The results are discussed in relation to possible effects of chronic stimulation by endogenous type C leukaemia virus upon differentiation of functional T cells or upon regulation by T cells of other T cell functions, including augmentation of antibody responses. ANSWER 6 OF 11 TOXCENTER COPYRIGHT 2003 ACS **DUPLICATE 5** AN 1981:44408 TOXCENTER CP Copyright 2003 BIOSIS DN BR20:48201 ΤI KINETIC ANALYSES OF NZB CYTO TOXIC LYMPHOCYTES ΑU HUSTON D P; CARMONA M A; STEINBERG A D BETHESDA, MD., USA. CS 44TH ANNUAL MEETING OF THE AMERICAN RHEUMATISM ASSOCIATION, ATLANTA, GA., SO USA, MAY 27-30, 1980. ARTHRITIS RHEUM. Arthritis Rheum., (1980) 23 (6), CODEN: ARHEAW. ISSN: 0004-3591. DTConference FS BIOSIS os BIOSIS 1981:48201 LΑ English Entered STN: 20011116 Last Updated on STN: 20011116 L3 ANSWER 7 OF 11 WPIDS (C) 2003 THOMSON DERWENT DUPLICATE 1 AN 2000-304694 [27] WPIDS DNC C2000-092673 Use of proteolytic enzymes to modulate hyperactive T cells, especially for symptomatic treatment of immune-mediated inflammatory diseases, e.g. multiple sclerosis, diabetes, arthritis or glomerulonephritis. DC B04 D16 IN RANSBERGER, K; STAUDER, G PA (MUCO-N) MUCOS PHARMA GMBH & CO CYC 21 PΙ DE 19847114 A1 20000420 (200027)\* WO 2000021547 A2 20000420 (200027) DE RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE W: CA US EP 1121146 A2 20010808 (200146) DE R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE ADT DE 19847114 A1 DE 1998-19847114 19981013; WO 2000021547 A2 WO 1999-EP7634 19991012; EP 1121146 A2 EP 1999-953778 19991012, WO 1999-EP7634 19991012 FDT EP 1121146 A2 Based on WO 200021547 PRAI DE 1998-19847114 19981013 2000-304694 [27] WPIDS AB DE 19847114 A UPAB: 20000606 NOVELTY - One or more proteolytic enzymes are used, optionally together with rutoside, to modulate hyperactive T cells

USE - For symptomatic treatment of immune-mediated inflammatory diseases, e.g. multiple sclerosis, type I diabetes, rheumatoid arthritis or glomerulonephritis.

AN

96:647818 PROMT

```
L3
     ANSWER 8 OF 11 WPIDS (C) 2003 THOMSON DERWENT
AN
     1999-061611 [06]
                        WPIDS
DNC C1999-018514
     Treatment of glomerulonephritis without side effects - using proteolytic
TT
     enzyme(s), especially trypsin, bromelain and/or papain, optionally in
     combination with rutoside.
DC
     B04 D16
     RANSBERGER, K; STAUDER, G
IN
      (MUCO-N) MUCOS PHARMA GMBH & CO
PΑ
CYC
                   A1 19981224 (199906) *
     DE 19726253
PΙ
     WO 9858667
                   A1 19981230 (199907) DE
        RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE
         W: US
     EP 920332
                   A1 19990609 (199927) DE
         R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE
     DE 19726253
                  C2 20000316 (200018)
ADT
     DE 19726253 A1 DE 1997-19726253 19970620; WO 9858667 A1 WO 1998-EP3769
     19980619; EP 920332 A1 EP 1998-934986 19980619, WO 1998-EP3769 19980619;
     DE 19726253 C2 DE 1997-19726253 19970620
     EP 920332 A1 Based on WO 9858667
PRAI DE 1997-19726253 19970620
     1999-061611 [06]
                       WPIDS
AB
     DE 19726253 A UPAB: 19990210
     Use of at least one proteolytic enzyme and optionally rutoside, for
     treating glomerulonephritis, is new.
          USE - The proteolytic enzymes may act by breaking up deposits of
     immune complexes in tissue affected by glomerulonephritis. They may also
     act by effecting altered cellular expression of cytokines, cytokine
     receptors, endogenous tissue metalloproteases or cellular adhesion
     molecules. Alternatively they may act by effecting reductions in the
     amount of hyperactive T-cells.
          ADVANTAGE - No damaging side effects are observed, even with use of
     the enzymes (or combination of enzymes) over a long period of time.
     Dwg.0/5
      ANSWER 9 OF 11 PHARMAML COPYRIGHT 2003 MARKETLETTER
1.3
ΑN
      1635213
               PHARMAML
TΙ
      Xoma Receives $8m From Genentech For Psoriasis Drug
SO
      Marketletter December 16, 1996
      Newsletter
DT
WC
      71
ТX
      - Xoma has received an $8.5 million milestone payment from Genentech to
      cover the expense of developing its psoriasis drug, hull24 (antiCD11a
      monoclonal antibody), through 1997. The company has started a Phase I
      trial of the product in 30-40 patients with moderate to severe psoriasis.
      The product is designed to block the hyperactive T
      cell reaction which occurs in psoriasis patients. It is also in
      development to treat organ transplant rejection.
L3
     ANSWER 10 OF 11 PHIN COPYRIGHT 2003 PJB
     1998:19358 PHIN
AN
DN
     B00600446
DED
     1 Oct 1998
     Rigel Inc.: A Laboratory in a Cell
     Bioventure-View (1998) No. 1310 pl
DT
     Newsletter
FS
     FULL
     ANSWER 11 OF 11 PROMT COPYRIGHT 2003 Gale Group
L3
```

TI Xoma Receives \$8m From Genentech For Psoriasis Drug

SO Marketletter, (16 Dec 1996) pp. N/A.

ISSN: 0951-3175.

LA English

WC 71

\*FULL TEXT IS AVAILABLE IN THE ALL FORMAT\*

- Xoma has received an \$8.5 million milestone payment from Genentech to cover the expense of developing its psoriasis drug, hull24 (antiCD11a monoclonal antibody), through 1997. The company has started a Phase I trial of the product in 30-40 patients with moderate to severe psoriasis. The product is designed to block the hyperactive T cell reaction which occurs in psoriasis patients. It is also in development to treat organ transplant rejection.

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=> log y		
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	ENTRY	SESSION
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